

Table 2 Body weights
Male, Female

Sex	Group and dose	Body weight (g) on day													
		1	8	15	22	29	36	43	50	57	64	71	78	85	
Male	Control	N	20	20	20	20	20	20	20	20	20	20	20	20	20
		Mean	212.9	276.8	333.8	378.3	426.4	465.9	495.0	526.4	548.1	566.7	585.6	601.7	615.6
		S.D.	±8.9	±14.2	±18.2	±22.3	±26.5	±29.5	±33.2	±39.4	±40.7	±42.5	±46.8	±49.7	±49.9
	4 mg/kg	N	20	20	20	20	20	20	20	20	20	20	20	20	20
		Mean	211.7	273.4	325.9	367.7	412.5	449.1	475.6	504.5	524.0	542.1	559.0	574.5	590.4
		S.D.	±9.0	±14.9	±21.7	±27.1	±33.4	±40.2	±47.6	±53.3	±55.1	±57.9	±59.8	±61.3	±63.6
	20 mg/kg	N	20	20	20	20	20	20	20	20	20	20	20	20	20
		Mean	211.2	277.0	333.4	376.8	421.3	458.5	487.3	514.8	537.1	557.7	576.2	596.2	610.6
		S.D.	±9.7	±17.4	±25.2	±29.8	±34.6	±40.0	±44.4	±47.7	±49.3	±52.9	±56.2	±59.9	±62.0
	100 mg/kg	N	20	20	20	20	20	20	20	20	20	20	20	20	20
		Mean	210.4	281.7	342.1	389.9	439.0	478.1	506.9	536.8	560.2	580.3	601.8	620.5	630.5
		S.D.	±8.8	±15.4	±23.0	±28.7	±32.5	±36.9	±38.8	±43.4	±44.4	±46.8	±50.3	±52.6	±56.0
Female	Control	N	20	20	20	20	20	20	20	20	20	20	20	20	
		Mean	157.2	186.0	215.5	233.2	252.7	263.2	277.1	286.1	294.5	304.1	309.2	316.3	321.0
		S.D.	±5.9	±8.8	±11.4	±13.7	±17.1	±17.3	±18.5	±20.5	±23.4	±23.2	±25.8	±26.3	±28.9
	4 mg/kg	N	20	20	20	20	20	20	20	20	20	20	20	20	20
		Mean	157.3	186.3	215.4	233.8	250.0	263.3	274.2	283.4	290.7	300.3	305.4	311.1	315.8
		S.D.	±7.4	±10.5	±12.4	±12.3	±14.4	±15.4	±15.8	±14.8	±17.2	±17.3	±19.4	±19.6	±20.4
	20 mg/kg	N	20	20	20	20	20	20	20	20	20	20	20	20	20
		Mean	156.7	186.7	217.4	233.2	251.7	264.7	274.9	284.5	291.8	301.4	309.3	315.5	319.5
		S.D.	±7.6	±9.8	±13.9	±14.9	±15.7	±15.9	±19.7	±20.0	±21.8	±22.7	±25.2	±25.0	±26.0
	100 mg/kg	N	20	20	20	20	20	20	20	20	20	20	20	20	20
		Mean	156.8	189.1	218.6	237.0	254.3	266.4	279.6	289.2	296.8	303.9	309.9	317.3	321.3
		S.D.	±8.6	±10.9	±17.0	±17.6	±22.3	±21.3	±23.9	±25.4	±27.0	±27.1	±27.8	±29.7	±32.6

Not significantly different from control.

Table 2 - continued
Body weights
Male, Female

Sex	Group and dose	Body weight (g) on day												
		91	92	120	148	176	204	232	260	288	316	344	364	
Male	Control	N	20	10	10	10	10	10	10	10	10	10	10	10
		Mean	624.3	615.4	659.2	692.9	729.8	752.6	779.8	814.6	839.3	849.8	866.0	876.1
		S.D.	±49.1	±51.1	±59.5	±65.2	±71.2	±73.5	±77.0	±88.4	±94.7	±101.8	±103.8	±107.7
	4 mg/kg	N	19	10	9	9	9	9	9	9	9	9	8	8
		Mean	602.2	594.0	635.7	671.9	705.7	727.1	752.1	781.1	796.5	812.3	813.4	823.7
		S.D.	±66.3	±69.1	±70.0	±71.2	±80.3	±78.6	±79.7	±85.4	±88.5	±97.0	±81.8	±82.1
	20 mg/kg	N	20	10	10	10	10	9	9	8	8	8	8	8
		Mean	621.8	625.6	669.9	706.5	737.9	778.1	800.4	802.1	823.7	836.4	856.0	866.8
		S.D.	±65.7	±84.0	±96.1	±105.5	±110.8	±119.2	±126.9	±133.2	±135.2	±139.0	±140.4	±140.4
	100 mg/kg	N	19	10	10	10	10	10	10	10	10	10	10	10
		Mean	643.5	635.1	680.7	716.2	750.1	786.7	816.0	844.0	866.0	880.2	901.2	907.3
		S.D.	±57.7	±72.6	±81.9	±86.8	±96.7	±107.9	±118.8	±130.9	±135.4	±139.1	±139.8	±136.5
Female	Control	N	20	10	10	10	10	10	10	10	10	10	10	
		Mean	323.7	314.2	326.3	338.8	353.8	367.5	385.0	399.1	414.8	428.6	441.0	449.6
		S.D.	±28.2	±30.5	±32.2	±33.4	±40.8	±49.4	±53.2	±56.2	±59.6	±69.4	±70.9	±78.0
	4 mg/kg	N	20	10	10	10	10	10	10	10	10	10	10	10
		Mean	320.1	315.0	329.0	340.5	352.9	368.6	385.1	391.3	408.5	414.4	423.4	426.0
		S.D.	±20.5	±20.9	±21.5	±24.8	±25.9	±27.2	±28.4	±36.6	±42.9	±45.4	±48.4	±54.3
	20 mg/kg	N	20	10	10	10	10	10	10	10	10	9	9	9
		Mean	323.9	314.0	325.5	340.6	357.2	372.2	388.2	401.4	418.5	425.6	445.0	454.1
		S.D.	±27.7	±11.4	±9.2	±15.6	±17.1	±19.1	±23.2	±23.7	±26.5	±32.9	±33.7	±39.4
	100 mg/kg	N	20	10	10	10	10	10	10	10	10	10	10	10
		Mean	326.8	329.1	346.0	358.2	374.6	388.2	408.5	421.7	439.3	453.5	467.4	471.0
		S.D.	±30.6	±34.7	±39.5	±48.0	±51.3	±56.3	±67.1	±69.0	±78.1	±84.0	±96.4	±94.1

Not significantly different from control.

Table 3 Food consumption
Male, Female

Sex	Group and dose		Food consumption(g) on day												
			2	8	15	22	29	36	43	50	57	64	71	78	85
Male	Control	N	20	20	20	20	20	20	20	20	20	20	20	20	20
		Mean	23.3	25.1	25.2	25.6	26.0	25.5	24.2	25.1	23.4	23.5	22.9	22.9	21.9
		S.D.	±1.6	±2.0	±2.7	±2.2	±2.5	±2.6	±2.0	±3.1	±2.4	±2.7	±3.1	±3.0	±2.4
	4 mg/kg	N	20	20	20	20	20	20	20	20	20	20	20	20	20
		Mean	23.0	23.9	24.6	25.0	24.5	25.0	23.3	24.0	23.2	22.9	21.9	21.8	22.8
		S.D.	±1.6	±2.3	±3.1	±2.8	±3.8	±2.9	±2.8	±3.4	±3.1	±2.8	±3.2	±3.0	±4.0
	20 mg/kg	N	20	20	20	20	20	20	20	20	20	20	20	20	20
		Mean	24.1	25.1	25.5	26.2	25.4	25.7	25.0	25.7	24.7	24.6	24.0	23.5	23.5
		S.D.	±2.6	±2.8	±3.0	±3.2	±3.4	±3.0	±3.3	±2.8	±3.1	±3.0	±3.4	±3.3	±3.2
	100 mg/kg	N	20	20	20	20	20	20	20	20	20	20	20	20	20
		Mean	24.1	24.6	26.5	26.8	26.3	25.9	25.2	25.8	26.1*	24.5	24.7	24.6	23.8*
		S.D.	±1.7	±2.6	±2.8	±4.0	±3.4	±3.3	±2.4	±2.6	±3.2	±3.3	±3.9	±3.4	±6.2
Female	Control	N	20	20	20	20	20	20	20	20	20	20	20	20	
		Mean	16.0	15.4	16.4	16.4	17.2	15.6	17.5	16.0	16.5	15.7	15.2	16.5	14.5
		S.D.	±1.7	±1.9	±2.3	±2.3	±2.1	±2.1	±2.1	±2.0	±2.7	±2.2	±2.2	±2.3	±2.3
	4 mg/kg	N	20	20	20	20	20	20	20	20	20	20	20	20	20
		Mean	16.2	16.0	16.4	16.4	16.6	16.4	16.7	15.3	15.9	15.8	16.3	14.9	15.1
		S.D.	±1.5	±1.9	±2.3	±1.7	±3.0	±2.4	±3.0	±2.4	±2.7	±1.6	±2.3	±2.7	±2.5
	20 mg/kg	N	20	20	20	20	20	20	20	20	20	20	20	20	20
		Mean	16.2	15.2	16.6	16.2	16.9	16.7	16.5	16.9	16.4	15.9	16.4	16.7	15.6
		S.D.	±2.2	±1.9	±2.3	±2.7	±3.0	±2.5	±2.5	±2.8	±3.0	±2.7	±2.8	±3.1	±3.1
	100 mg/kg	N	20	20	20	20	20	20	20	20	20	20	20	20	20
		Mean	16.5	15.7	16.9	16.9	18.3	16.6	17.2	17.1	17.5	15.5	16.1	17.3	15.7
		S.D.	±1.2	±1.3	±2.5	±2.7	±3.7	±2.7	±2.7	±2.4	±2.6	±2.4	±3.2	±2.9	±2.9

*: P<0.05 (significantly different from control).

Table 3 - continued Food consumption
Male, Female

Sex	Group and dose	Food consumption(g) on day												
		91	92	120	148	176	204	232	260	288	316	344	364	
Male	Control	N	20	10	10	10	10	10	10	10	10	10	10	10
		Mean	22.8	21.7	22.0	21.4	20.3	18.6	18.8	20.1	16.9	17.6	18.6	17.3
		S.D.	±2.0	±1.4	±2.5	±1.8	±2.7	±2.2	±2.4	±2.4	±2.2	±1.6	±2.9	±1.9
	4 mg/kg	N	19	10	9	9	9	9	9	9	9	9	8	8
		Mean	22.3	21.6	21.3	21.3	20.4	19.0	18.7	19.4	17.6	18.1	17.8	16.6
		S.D.	±3.3	±4.4	±3.6	±3.3	±4.2	±2.3	±3.0	±4.4	±3.9	±2.7	±1.6	±2.5
	20 mg/kg	N	20	10	10	10	10	9	9	8	8	8	8	8
		Mean	24.1	23.1	23.5	22.2	21.2	19.9	18.7	19.1	17.0	18.7	18.5	17.2
		S.D.	±3.7	±3.5	±3.3	±3.0	±2.4	±3.4	±3.9	±4.3	±4.0	±2.5	±2.8	±3.5
	100 mg/kg	N	19	10	10	10	10	10	10	10	10	10	10	10
		Mean	24.9	21.8	23.4	23.1	22.1	20.5	19.5	19.6	19.9	19.9	19.1	17.3
		S.D.	±3.4	±3.3	±3.2	±3.3	±2.6	±4.1	±3.7	±3.6	±2.8	±3.5	±1.9	±1.9
Female	Control	N	20	10	10	10	10	10	10	10	10	10	10	10
		Mean	15.8	14.9	14.7	15.5	15.5	13.7	13.7	14.3	15.4	16.1	12.9	13.0
		S.D.	±2.7	±1.2	±2.2	±1.6	±1.9	±2.8	±0.9	±1.5	±1.7	±2.6	±2.6	±2.6
	4 mg/kg	N	20	10	10	10	10	10	10	10	10	10	10	10
		Mean	16.8	15.7	16.4	15.5	15.9	15.0	14.0	13.0	15.5	15.4	13.8	13.3
		S.D.	±2.5	±3.5	±2.4	±2.3	±2.6	±2.4	±2.5	±2.7	±2.0	±1.3	±1.7	±4.1
	20 mg/kg	N	20	10	10	10	10	10	10	10	10	9	9	9
		Mean	16.5	14.5	15.1	16.3	16.6	15.0	13.8	14.3	15.6	15.9	14.2	15.3
		S.D.	±2.8	±1.5	±1.9	±2.8	±2.4	±2.9	±3.2	±2.7	±2.5	±3.4	±2.6	±3.0
	100 mg/kg	N	20	10	10	10	10	10	10	10	10	10	10	10
		Mean	17.3	15.9	15.8	17.0	18.0	15.6	15.4	14.3	16.4	15.4	15.0	15.1
		S.D.	±2.5	±1.7	±2.7	±2.4	±2.4	±2.5	±2.7	±2.7	±2.3	±3.8	±3.4	±2.6

Not significantly different from control.

Table 4 Urinary findings
Male, Female, 13w

Sex	Group and dose		Urine volume (mL/24hr)	Osmotic pressure (Osm/kg)	Specific gravity	Na (mEq/24hr)	K (mEq/24hr)	Cl (mEq/24hr)
Male	Control	N	10	10	10	10	10	10
		Mean	13.4	1.512	1.048	0.944	2.187	1.256
		S. D.	±5.1	±0.439	±0.015	±0.273	±0.502	±0.401
	4 mg/kg	N	9	9	9	9	9	9
		Mean	17.6	1.452	1.048	0.852	2.145	1.250
		S. D.	±14.1	±0.596	±0.023	±0.416	±0.777	±0.595
	20 mg/kg	N	10	10	10	10	10	10
		Mean	15.4	1.506	1.047	0.988	2.433	1.422
		S. D.	±7.5	±0.433	±0.015	±0.297	±0.387	±0.387
	100 mg/kg	N	9	9	9	9	9	9
		Mean	14.5	1.454	1.047	1.027	2.449	1.488
		S. D.	±4.7	±0.294	±0.009	±0.456	±0.440	±0.357
Female	Control	N	10	10	10	10	10	10
		Mean	12.5	1.200	1.039	0.755	1.679	1.101
		S. D.	±7.6	±0.479	±0.017	±0.359	±0.490	±0.490
	4 mg/kg	N	10	10	10	10	10	10
		Mean	12.8	1.206	1.040	0.919	1.846	1.267
		S. D.	±7.1	±0.410	±0.014	±0.240	±0.507	±0.318
	20 mg/kg	N	10	10	10	10	10	10
		Mean	8.8	1.617	1.054	0.774	1.908	1.204
		S. D.	±3.3	±0.385	±0.013	±0.300	±0.463	±0.315
	100 mg/kg	N	10	10	10	10	10	10
		Mean	12.3	1.169	1.039	0.883	1.807	1.231
		S. D.	±5.3	±0.400	±0.014	±0.232	±0.368	±0.196

Not significantly different from control.

One male in the 4 mg/kg group was imminently sacrificed when moribund and one male in the 100 mg/kg group died.

Table 4 - continued

Urinary findings
Male, Female, 13w

Sex	Group and dose	Number of animals	Color		pH						Protein			Glucose	Ketone body
			PY	Y	5.5	6.0	6.5	7.0	8.0	8.5	-	±	+	-	-
Male	Control	10	0	10	0	0	0	1	8	1	5	3	2	10	10
	4 mg/kg	9	1	8	0	0	0	1	6	2	2	4	3	9	9
	20 mg/kg	10	1	9	0	0	0	0	8	2	4	4	2	10	10
	100 mg/kg	9	0	9	0	0	0	2	5	2	0	5	4	9	9
Female	Control	10	1	9	0	2	0	4	3	1	8	2	0	10	10
	4 mg/kg	10	1	9	1	0	0	0	7	2	10	0	0	10	10
	20 mg/kg	10	0	10	0	0	1	0	8	1	9	1	0	10	10
	100 mg/kg	10	0	10	0	3	0	1	4	2	10	0	0	10	10

Not significantly different from control.

Abbreviation: PY, pale yellow; Y, yellow; YB, yellowish brown; B, brown.

Grade sign: -, none; ±, trace; +, slight; ++, moderate; +++, severe; +++++, very severe.

One male in the 4 mg/kg group was imminently sacrificed when moribund and one male in the 100 mg/kg group died.

Table 4 - continued
Urinary findings
Male, Female, 13w

Sex	Group and dose	Number of animals	Urinary findings		
			Bilirubin	Occult blood	Urobilinogen (mg/dL)
			-	-	<1
Male	Control	10	10	10	10
	4 mg/kg	9	9	9	9
	20 mg/kg	10	10	10	10
	100 mg/kg	9	9	9	9
Female	Control	10	10	10	10
	4 mg/kg	10	10	10	10
	20 mg/kg	10	10	10	10
	100 mg/kg	10	10	10	10

Not significantly different from control.

Grade sign: -, none; ±, trace; +, slight; ++, moderate; +++, severe; +++++, very severe.

One male in the 4 mg/kg group was imminently sacrificed when moribund and one male in the 100 mg/kg group died.

Table 4 - continued
Urinary findings
Male, Female, 13w

Sex	Group and dose	Number of animals	Urinary sediment					Crystals		
			Epithelial cells	Erythrocytes	Leukocytes	Casts	-	+	++	
			-	-	-	-	-	+	++	
Male	Control	10	10	10	10	10	9	1	0	
	4 mg/kg	9	9	9	9	9	7	2	0	
	20 mg/kg	10	10	10	10	10	8	2	0	
	100 mg/kg	9	9	9	9	9	8	0	1	
Female	Control	10	10	10	10	10	10	0	0	
	4 mg/kg	10	10	10	10	10	10	0	0	
	20 mg/kg	10	10	10	10	10	10	0	0	
	100 mg/kg	10	10	10	10	10	10	0	0	

Not significantly different from control.

Grade signs are as follows.

Epithelial cells: -, < 3/field; +, 3/field \leq and < 10/field; ++, 10/field \leq and < 20/field; +++, \geq 20/field.

Erythrocytes: -, < 10/field; +, 10/field \leq and < 30/field; ++, 30/field \leq and < 100/field; +++, countless.

Leukocytes: -, < 3/field; +, 3/field \leq and < 20/field; ++, 20/field \leq and < 40/field; +++, \geq 40/field.

Casts: -, none; +, \geq 1/all field.

Crystals: -, < 10/field; +, 10/field \leq and < 20/field; ++, 20/field \leq and < 30/field; +++, countless.

One male in the 4 mg/kg group was imminently sacrificed when moribund and one male in the 100 mg/kg group died.

Table 5 Urinary findings
Male, Female, 52w

Sex	Group and dose		Urine volume (mL/24hr)	Osmotic pressure (Osm/kg)	Specific gravity	Na (mEq/24hr)	K (mEq/24hr)	Cl (mEq/24hr)
Male	Control	N	10	10	10	10	10	10
		Mean	8.7	1.714	1.058	0.506	1.594	0.684
		S.D.	±3.4	±0.415	±0.019	±0.203	±0.380	±0.309
	4 mg/kg	N	8	8	8	8	8	8
		Mean	10.4	1.435	1.050	0.501	1.395	0.652
		S.D.	±5.7	±0.527	±0.019	±0.183	±0.232	±0.285
	20 mg/kg	N	8	8	8	8	8	8
		Mean	11.3	1.388	1.048	0.611	1.583	0.834
		S.D.	±5.9	±0.446	±0.017	±0.457	±0.560	±0.569
	100 mg/kg	N	10	10	10	10	10	10
		Mean	13.0	1.275	1.044	0.701	1.648	0.832
		S.D.	±4.8	±0.259	±0.011	±0.356	±0.246	±0.365
Female	Control	N	10	10	10	10	10	10
		Mean	14.2	0.992	1.034	0.844	1.831	1.134
		S.D.	±5.7	±0.280	±0.010	±0.254	±0.377	±0.301
	4 mg/kg	N	10	10	10	10	10	10
		Mean	15.1	1.046	1.037	0.891	1.930	1.193
		S.D.	±7.3	±0.469	±0.019	±0.281	±0.372	±0.388
	20 mg/kg	N	9	9	9	9	9	9
		Mean	18.0	0.810	1.028	0.879	1.971	1.229
		S.D.	±5.7	±0.256	±0.009	±0.278	±0.410	±0.369
	100 mg/kg	N	10	10	10	10	10	10
		Mean	13.8	1.158	1.042	0.834	1.927	1.152
		S.D.	±7.1	±0.417	±0.017	±0.244	±0.416	±0.315

Not significantly different from control.

One male in the 4 mg/kg group died and one male in the 4 mg/kg group was imminently sacrificed when moribund.

Two males and one female in the 20 mg/kg group died.

Table 5 - continued
Urinary findings
Male, Female, 52w

Sex	Group and dose	Number of animals	Color		pH								Protein					Glucose	
			PY	Y	5.0	5.5	6.0	6.5	7.0	7.5	8.0	8.5	-	±	+	++	+++	-	
Male	Control	10	0	10	0	0	0	0	0	0	2	7	1	0	1	6	2	1	10
	4 mg/kg	8	0	8	0	0	0	0	0	0	3	4	1	0	1	5	1	1	8
	20 mg/kg	8	0	8	0	0	0	1	2	3	2	0*	0	0	3	4	1	8	
	100 mg/kg	10	0	10	0	0	0	0	5	1	4	0	0	2	1	5	2	10	
Female	Control	10	0	10	1	1	0	1	3	0	4	0	4	3	1	2	0	10	
	4 mg/kg	10	2	8	0	2	0	1	3	1	3	0	2	4	2	2	0	10	
	20 mg/kg	9	2	7	2	1	2	2	2	0	0	0	2	1	2	4	0	9	
	100 mg/kg	10	2	8	0	2	0	0	4	0	4	0	2	1	3	1	3	10	

*: P<0.05 (significantly different from control).

Abbreviation: PY, pale yellow; Y, yellow; YB, yellowish brown; B, brown.

Grade sign: -, none; ±, trace; +, slight; ++, moderate; +++, severe; +++++, very severe.

One male in the 4 mg/kg group died and one male in the 4 mg/kg group was imminently sacrificed when moribund.

Two males and one female in the 20 mg/kg group died.

Table 5 -- continued

Urinary findings
Male, Female, 52w

Sex	Group and dose	Number of animals	Ketone body	Bilirubin	Occult blood	Urobilinogen (mg/dL)
			-	-	-	<1
Male	Control	10	10	10	10	10
	4 mg/kg	8	8	8	8	8
	20 mg/kg	8	8	8	8	8
	100 mg/kg	10	10	10	10	10
Female	Control	10	10	10	10	10
	4 mg/kg	10	10	10	10	10
	20 mg/kg	9	9	9	9	9
	100 mg/kg	10	10	10	10	10

Not significantly different from control.

Grade sign: -, none; ±, trace; +, slight; ++, moderate; +++, severe; +++++, very severe.

One male in the 4 mg/kg group died and one male in the 4 mg/kg group was imminently sacrificed when moribund.

Two males and one female in the 20 mg/kg group died.

Table 5 - continued
Urinary findings
Male, Female, 52w

Sex	Group and dose	Number of animals	Urinary sediment				
			Epithelial cells	Erythrocytes	Leukocytes	Casts	Crystals
			-	-	- + ++	-	-
Male	Control	10	10	10	9 1 0	10	10
	4 mg/kg	8	8	8	6 2 0	8	8
	20 mg/kg	8	8	8	6 2 0	8	8
	100 mg/kg	10	10	10	9 1 0	10	10
Female	Control	10	10	10	9 0 1	10	10
	4 mg/kg	10	10	10	10 0 0	10	10
	20 mg/kg	9	9	9	9 0 0	9	9
	100 mg/kg	10	10	10	10 0 0	10	10

Not significantly different from control.

Grade signs are as follows.

Epithelial cells: -, < 3/field; +, 3/field \leq and < 10/field; ++, 10/field \leq and < 20/field; +++, \geq 20/field.
 Erythrocytes : -, < 10/field; +, 10/field \leq and < 30/field; ++, 30/field \leq and < 100/field; +++, countless.
 Leukocytes : -, < 3/field; +, 3/field \leq and < 20/field; ++, 20/field \leq and < 40/field; +++, \geq 40/field.
 Casts : -, none; +, \geq 1/all field.
 Crystals : -, < 10/field; +, 10/field \leq and < 20/field; ++, 20/field \leq and < 30/field; +++, countless.

One male in the 4 mg/kg group died and one male in the 4 mg/kg group was imminently sacrificed when moribund.
 Two males and one female in the 20 mg/kg group died.

Table 6 Hematological findings
Male, Female, 13w

Sex	Group and dose		Leukocytes (10 ³ / μ L)	Erythrocytes (10 ⁴ / μ L)	Hemoglobin (g/dL)	Hematocrit (%)	MCV (fL)	MCH (pg)	MCHC (g/dL)	Reticulocyte (10 ⁴ / μ L)	Platelets (10 ⁴ / μ L)
Male	Control	N	10	10	10	10	10	10	10	10	10
		Mean	10.08	853	14.5	42.8	50.1	17.0	34.0	17.7	112.6
		S.D.	± 3.06	± 35	± 0.6	± 1.9	± 0.6	± 0.2	± 0.5	± 2.3	± 15.8
	4 mg/kg	N	9	9	9	9	9	9	9	9	9
		Mean	9.09	863	14.6	42.8	49.7	16.9	34.0	17.6	109.0
		S.D.	± 2.08	± 33	± 0.4	± 1.3	± 1.2	± 0.7	± 0.7	± 2.5	± 12.4
	20 mg/kg	N	10	10	10	10	10	10	10	10	10
		Mean	10.35	846	14.5	42.1	49.9	17.2	34.4	19.5	113.6
		S.D.	± 1.88	± 32	± 0.6	± 1.8	± 1.5	± 0.5	± 0.7	± 3.2	± 10.9
	100 mg/kg	N	9	9	9	9	9	9	9	9	9
		Mean	9.28	857	14.4	42.0	49.1	16.8	34.2	18.0	111.6
		S.D.	± 2.21	± 38	± 0.3	± 1.1	± 1.7	± 0.7	± 0.5	± 2.4	± 11.3
Female	Control	N	10	10	10	10	10	10	10	10	10
		Mean	4.61	789	14.1	40.3	51.1	17.9	35.0	14.5	115.1
		S.D.	± 1.47	± 48	± 0.5	± 1.7	± 1.7	± 0.7	± 0.5	± 4.1	± 13.9
	4 mg/kg	N	10	10	10	10	10	10	10	10	10
		Mean	4.65	784	14.1	40.0	51.1	18.0	35.2	13.3	119.8
		S.D.	± 1.08	± 37	± 0.3	± 0.8	± 1.8	± 0.7	± 0.4	± 2.2	± 11.4
	20 mg/kg	N	10	10	10	10	10	10	10	10	10
		Mean	5.84	772	13.7	39.4	51.1	17.8	34.8	15.5	115.3
		S.D.	± 0.67	± 40	± 0.6	± 1.6	± 1.4	± 0.5	± 0.5	± 2.8	± 14.5
	100 mg/kg	N	10	10	10	10	10	10	10	10	10
		Mean	5.53	778	13.9	39.1	50.3	17.9	35.5	14.6	120.1
		S.D.	± 1.57	± 28	± 0.6	± 1.5	± 1.4	± 0.6	± 0.7	± 2.5	± 13.1

Not significantly different from control.

One male in the 4 mg/kg group was imminently sacrificed when moribund and one male in the 100 mg/kg group died.

Table 6 - continued
Hematological findings
Male, Female, 13w

Sex	Group and dose		PT (sec)	APTT (sec)
Male	Control	N	10	10
		Mean	13.8	22.7
		S.D.	±1.3	±1.9
	4 mg/kg	N	9	9
		Mean	15.2	24.6
		S.D.	±2.3	±2.6
	20 mg/kg	N	10	10
		Mean	14.4	22.5
		S.D.	±1.3	±2.0
	100 mg/kg	N	9	9
		Mean	14.7	24.4
		S.D.	±0.7	±2.3
Female	Control	N	10	10
		Mean	12.7	19.1
		S.D.	±0.6	±0.7
	4 mg/kg	N	10	10
		Mean	12.5	19.3
		S.D.	±0.6	±1.1
	20 mg/kg	N	10	10
		Mean	12.5	19.5
		S.D.	±0.4	±0.8
	100 mg/kg	N	10	10
		Mean	12.4	20.1
		S.D.	±0.5	±1.3

Not significantly different from control.

One male in the 4 mg/kg group was imminently sacrificed when moribund and one male in the 100 mg/kg group died.

Table 6 - continued Hematological findings
Male, Female, 13w

Sex	Group and dose		Differential leukocyte count					
			Eosinophils	Neutrophils	Lymphocytes	Basophils	Monocytes	Large unstained cells
			(10 ² / μ L)	(10 ² / μ L)	(10 ² / μ L)	(10 ² / μ L)	(10 ² / μ L)	(10 ² / μ L)
Male	Control	N	10	10	10	10	10	10
		Mean	1.4	18.6	77.9	0.3	1.8	0.8
		S. D.	± 0.5	± 8.1	± 29.8	± 0.2	± 0.8	± 0.6
	4 mg/kg	N	9	9	9	9	9	9
		Mean	1.9	15.9	70.5	0.2	1.6	0.8
		S. D.	± 0.4	± 5.9	± 18.3	± 0.1	± 0.5	± 0.6
	20 mg/kg	N	10	10	10	10	10	10
		Mean	1.5	20.6	78.3	0.3	2.0	0.9
		S. D.	± 0.6	± 9.5	± 13.9	± 0.1	± 0.7	± 0.4
	100 mg/kg	N	9	9	9	9	9	9
		Mean	1.7	18.8	69.5	0.3	1.7	0.8
		S. D.	± 0.5	± 5.1	± 19.3	± 0.2	± 0.5	± 0.4
Female	Control	N	10	10	10	10	10	10
		Mean	0.8	5.7	38.6	0.1	0.7	0.3
		S. D.	± 0.3	± 1.4	± 13.2	± 0.1	± 0.3	± 0.2
	4 mg/kg	N	10	10	10	10	10	10
		Mean	0.7	6.1	38.4	0.1	0.7	0.4
		S. D.	± 0.2	± 2.7	± 9.0	± 0.0	± 0.2	± 0.3
	20 mg/kg	N	10	10	10	10	10	10
		Mean	0.7	6.5	49.7	0.1	0.8	0.5
		S. D.	± 0.2	± 2.3	± 6.5	± 0.0	± 0.4	± 0.2
	100 mg/kg	N	10	10	10	10	10	10
		Mean	0.9	6.0	47.0	0.1	0.8	0.6*
		S. D.	± 0.3	± 1.5	± 15.1	± 0.1	± 0.4	± 0.3

*: P<0.05 (significantly different from control).

One male in the 4 mg/kg group was imminently sacrificed when moribund and one male in the 100 mg/kg group died.

Table 7 Hematological findings
Male, Female, 52w

Sex	Group and dose		Leukocytes (10 ³ / μ L)	Erythrocytes (10 ⁴ / μ L)	Hemoglobin (g/dL)	Hematocrit (%)	MCV (fL)	MCH (pg)	MCHC (g/dL)	Reticulocyte (10 ⁴ / μ L)	Platelets (10 ⁴ / μ L)	
Male	Control	N	10	10	10	10	10	10	10	10	10	
		Mean	7.21	840	14.0	45.0	53.6	16.7	31.2	15.6	103.4	
		S.D.	± 1.47	± 47	± 0.9	± 2.9	± 2.2	± 0.6	± 0.5	± 3.9	± 8.3	
	4 mg/kg	N	8	8	8	8	8	8	8	8	8	8
		Mean	7.42	843	14.2	45.2	53.7	16.8	31.4	16.1	103.4	
		S.D.	± 1.08	± 67	± 1.1	± 3.0	± 2.3	± 0.6	± 0.5	± 4.2	± 14.2	
	20 mg/kg	N	8	8	8	8	8	8	8	8	8	8
		Mean	7.51	843	13.6	43.6	51.6	16.1	31.2	17.5	113.3	
		S.D.	± 2.15	± 91	± 2.3	± 6.1	± 3.9	± 1.8	± 1.4	± 10.1	± 33.6	
	100 mg/kg	N	10	10	10	10	10	10	10	10	10	10
		Mean	11.94**	881	13.4	43.1	48.9**	15.2*	31.0	17.6	110.5	
		S.D.	± 2.51	± 45	± 0.8	± 2.3	± 3.0	± 1.0	± 0.6	± 2.5	± 10.5	
Female	Control	N	10	10	10	10	10	10	10	10	10	
		Mean	4.75	747	13.9	41.8	56.0	18.6	33.2	14.4	98.8	
		S.D.	± 0.80	± 39	± 0.5	± 1.7	± 2.6	± 0.7	± 1.0	± 3.7	± 15.1	
	4 mg/kg	N	10	10	10	10	10	10	10	10	10	10
		Mean	7.10	657	12.2	37.3*	57.2	18.7	32.7	17.6	97.3	
		S.D.	± 6.88	± 117	± 2.0	± 5.2	± 3.6	± 0.5	± 1.3	± 9.2	± 23.9	
	20 mg/kg	N	9	9	9	9	9	9	9	9	9	9
		Mean	4.24	676	12.7*	38.6	57.5	18.9	32.9	20.9	109.1	
		S.D.	± 1.17	± 99	± 1.6	± 4.1	± 4.3	± 1.0	± 0.9	± 18.6	± 23.7	
	100 mg/kg	N	10	10	10	10	10	10	10	10	10	10
		Mean	5.33	693	12.4**	37.7*	54.5	18.0	33.0	13.7	105.3	
		S.D.	± 1.78	± 61	± 0.9	± 3.2	± 3.4	± 1.2	± 0.5	± 5.2	± 10.9	

*: P<0.05, **: P<0.01 (significantly different from control).

One male in the 4 mg/kg group died and one male in the 4 mg/kg group was imminently sacrificed when moribund.
Two males and one female in the 20 mg/kg group died.

Table 7 - continued
Hematological findings
Male, Female, 52w

Sex	Group and dose		PT (sec)	APTT (sec)
Male	Control	N	10	10
		Mean	14.2	22.1
		S.D.	±1.2	±1.3
	4 mg/kg	N	8	8
		Mean	14.8	21.7
		S.D.	±1.4	±2.1
	20 mg/kg	N	8	8
		Mean	14.2	21.9
		S.D.	±1.4	±1.5
	100 mg/kg	N	10	10
		Mean	19.0**	25.5**
		S.D.	±3.0	±2.1
Female	Control	N	10	10
		Mean	12.6	18.1
		S.D.	±0.6	±1.9
	4 mg/kg	N	10	10
		Mean	12.0	17.7
		S.D.	±1.3	±1.9
	20 mg/kg	N	9	9
		Mean	12.2	18.2
		S.D.	±0.4	±0.8
	100 mg/kg	N	10	10
		Mean	11.6*	17.9
		S.D.	±0.7	±1.6

*: P<0.05, **: P<0.01 (significantly different from control).

One male in the 4 mg/kg group died and one male in the 4 mg/kg group was imminently sacrificed when moribund.

Two males and one female in the 20 mg/kg group died.

Table 7 - continued
Hematological findings
Male, Female, 52w

Sex	Group and dose		Differential leukocyte count					
			Eosinophils	Neutrophils	Lymphocytes	Basophils	Monocytes	Large unstained cells
			(10 ² / μ L)	(10 ² / μ L)	(10 ² / μ L)	(10 ² / μ L)	(10 ² / μ L)	(10 ² / μ L)
Male	Control	N	10	10	10	10	10	10
		Mean	1.3	22.6	44.8	0.1	2.7	0.7
		S.D.	± 0.2	± 10.3	± 7.3	± 0.0	± 0.9	± 0.3
	4 mg/kg	N	8	8	8	8	8	8
		Mean	1.2	21.3	48.4	0.1	2.2	0.9
		S.D.	± 0.3	± 8.2	± 8.5	± 0.1	± 0.4	± 0.3
	20 mg/kg	N	8	8	8	8	8	8
		Mean	1.1	25.5	44.9	0.1	2.6	0.9
		S.D.	± 0.5	± 14.5	± 16.1	± 0.1	± 0.6	± 0.6
	100 mg/kg	N	10	10	10	10	10	10
		Mean	1.4	41.8**	70.6**	0.2*	4.1**	1.5**
		S.D.	± 0.5	± 12.1	± 19.9	± 0.1	± 0.7	± 0.4
Female	Control	N	10	10	10	10	10	10
		Mean	0.8	14.8	29.9	0.1	1.6	0.4
		S.D.	± 0.2	± 5.5	± 6.0	± 0.1	± 0.5	± 0.2
	4 mg/kg	N	10	10	10	10	10	10
		Mean	0.8	32.3	33.6	0.1	2.7	1.6
		S.D.	± 0.3	± 44.1	± 20.1	± 0.2	± 2.5	± 3.0
	20 mg/kg	N	9	9	9	9	9	9
		Mean	0.8	13.3	26.2	0.0	1.7	0.5
		S.D.	± 0.3	± 5.1	± 8.5	± 0.0	± 0.5	± 0.1
	100 mg/kg	N	10	10	10	10	10	10
		Mean	0.7	13.5	36.0	0.1	2.1	0.9
		S.D.	± 0.3	± 7.3	± 12.6	± 0.1	± 0.8	± 0.8

*: P<0.05, **: P<0.01 (significantly different from control).

One male in the 4 mg/kg group died and one male in the 4 mg/kg group was imminently sacrificed when moribund.

Two males and one female in the 20 mg/kg group died.

Table 8 Biochemical findings
Male, Female, 13w

Sex	Group and dose		T. Protein (g/dL)	A/G ratio	α_1 -Globulin (%)	α_2 -Globulin (%)	β -Globulin (%)	γ -Globulin (%)	Albumin (%)	T. Bilirubin (mg/dL)	AST (IU/L)	ALT (IU/L)	
Male	Control	N	10	10	10	10	10	10	10	10	10	10	
		Mean	5.7	1.20	18.2	7.8	15.0	4.5	54.6	0.0	95	33	
		S.D.	±0.3	±0.08	±1.6	±0.6	±0.8	±0.6	±1.6	±0.0	±26	±25	
	4 mg/kg	N	9	9	9	9	9	9	9	9	9	9	9
		Mean	5.6	1.17	18.6	7.9	15.1	4.6	53.9	0.0	94	35	
		S.D.	±0.3	±0.06	±1.2	±0.5	±0.8	±0.8	±1.2	±0.0	±44	±31	
	20 mg/kg	N	10	10	10	10	10	10	10	10	10	10	10
		Mean	5.7	1.20	17.2	7.8	15.7	5.0	54.4	0.0	88	29	
		S.D.	±0.3	±0.13	±2.2	±0.4	±1.2	±1.3	±2.7	±0.0	±33	±30	
	100 mg/kg	N	9	9	9	9	9	9	9	9	9	9	9
		Mean	5.8	1.15	17.2	8.4	16.8**	4.0	53.5	0.0	74	26	
		S.D.	±0.3	±0.07	±1.4	±0.8	±1.0	±1.1	±1.5	±0.0	±7	±5	
Female	Control	N	10	10	10	10	10	10	10	10	10	10	
		Mean	6.3	1.74	14.0	5.8	12.0	4.8	63.5	0.1	78	24	
		S.D.	±0.3	±0.14	±1.2	±0.4	±0.9	±0.8	±1.9	±0.1	±14	±6	
	4 mg/kg	N	10	10	10	10	10	10	10	10	10	10	10
		Mean	6.4	1.73	13.9	6.0	12.2	4.6	63.3	0.0	90	22	
		S.D.	±0.3	±0.13	±0.5	±0.5	±0.5	±0.8	±1.9	±0.0	±17	±9	
	20 mg/kg	N	10	10	10	10	10	10	10	10	10	10	10
		Mean	6.5	1.78	13.6	5.7	12.5	4.3	63.9	0.0	105	32	
		S.D.	±0.2	±0.17	±1.8	±0.5	±0.9	±1.0	±2.3	±0.0	±69	±40	
	100 mg/kg	N	10	10	10	10	10	10	10	10	10	10	10
		Mean	6.4	1.51**	14.7	6.4	13.9**	4.8	60.2**	0.0*	73	18	
		S.D.	±0.3	±0.07	±2.0	±0.7	±0.6	±0.9	±1.1	±0.0	±20	±3	

*: P<0.05, **: P<0.01 (significantly different from control).

One male in the 4 mg/kg group was imminently sacrificed when moribund and one male in the 100 mg/kg group died.

Table 8 - continued
Biochemical findings
Male, Female, 13w

Sex	Group and dose		ALP (IU/L)	T.Cholesterol (mg/dL)	Triglycerides (mg/dL)	Phospholipids (mg/dL)	Glucose (mg/dL)	BUN (mg/dL)	Creatinine (mg/dL)	IP (mg/dL)	Ca (mg/dL)	Na (mEq/L)	
Male	Control	N	10	10	10	10	10	10	10	10	10	10	
		Mean	197	73	60	123	132	12.5	0.5	6.2	10.1	145.4	
		S.D.	±49	±16	±30	±23	±21	±1.1	±0.1	±0.5	±0.3	±0.8	
	4 mg/kg	N	9	9	9	9	9	9	9	9	9	9	9
		Mean	208	79	45	128	124	12.6	0.4	6.1	10.2	145.3	
		S.D.	±26	±25	±16	±34	±13	±1.9	±0.1	±0.6	±0.5	±0.7	
	20 mg/kg	N	10	10	10	10	10	10	10	10	10	10	10
		Mean	167	75	49	124	129	13.2	0.5	6.3	10.2	145.7	
		S.D.	±28	±10	±22	±14	±14	±1.0	±0.1	±0.6	±0.4	±1.1	
	100 mg/kg	N	9	9	9	9	9	9	9	9	9	9	9
		Mean	167	79	47	129	132	12.7	0.4	6.3	10.4	145.3	
		S.D.	±40	±9	±19	±16	±15	±1.7	±0.1	±0.5	±0.3	±0.6	
Female	Control	N	10	10	10	10	10	10	10	10	10	10	
		Mean	99	79	30	154	120	16.1	0.5	4.9	10.5	143.3	
		S.D.	±20	±13	±13	±23	±12	±3.0	±0.1	±1.2	±0.2	±0.8	
	4 mg/kg	N	10	10	10	10	10	10	10	10	10	10	10
		Mean	85	82	24	154	118	16.6	0.5	5.3	10.7	142.8	
		S.D.	±21	±16	±8	±25	±13	±2.6	±0.0	±0.9	±0.3	±0.8	
	20 mg/kg	N	10	10	10	10	10	10	10	10	10	10	10
		Mean	89	93*	22	172	123	15.1	0.5	5.4	10.8	142.9	
		S.D.	±32	±5	±11	±10	±8	±2.6	±0.1	±0.8	±0.3	±0.8	
	100 mg/kg	N	10	10	10	10	10	10	10	10	10	10	10
		Mean	88	99*	25	172	118	16.8	0.5	5.3	10.6	142.9	
		S.D.	±27	±12	±22	±19	±9	±3.5	±0.0	±1.1	±0.4	±0.8	

*: P<0.05 (significantly different from control).

One male in the 4 mg/kg group was imminently sacrificed when moribund and one male in the 100 mg/kg group died.

Table 8 - continued
Biochemical findings
Male, Female, 13w

Sex	Group and dose	Biochemical findings		
		K (mEq/L)	Cl (mEq/L)	
Male	Control	N	10	10
		Mean	4.40	104.5
		S.D.	±0.21	±1.1
	4 mg/kg	N	9	9
		Mean	4.29	104.1
		S.D.	±0.28	±2.0
	20 mg/kg	N	10	10
		Mean	4.38	104.3
		S.D.	±0.20	±1.1
	100 mg/kg	N	9	9
		Mean	4.49	102.6*
		S.D.	±0.16	±1.3
Female	Control	N	10	10
		Mean	4.01	105.3
		S.D.	±0.30	±0.8
	4 mg/kg	N	10	10
		Mean	4.02	104.6
		S.D.	±0.23	±1.3
	20 mg/kg	N	10	10
		Mean	4.02	104.7
		S.D.	±0.18	±0.8
	100 mg/kg	N	10	10
		Mean	4.05	104.2
		S.D.	±0.32	±1.9

*: P<0.05 (significantly different from control).

One male in the 4 mg/kg group was imminently sacrificed when moribund and one male in the 100 mg/kg group died.

Table 9 Biochemical findings
Male, Female, 52w

Sex	Group and dose		T.Protein (g/dL)	A/G ratio	α_1 -Globulin (%)	α_2 -Globulin (%)	β -Globulin (%)	γ -Globulin (%)	Albumin (%)	T.Bilirubin (mg/dL)	AST (IU/L)	ALT (IU/L)	
Male	Control	N	10	10	10	10	10	10	10	10	10	10	
		Mean	6.1	1.00	19.6	7.7	17.1	5.8	49.8	0.0	90	37	
		S.D.	±0.3	±0.12	±2.7	±1.2	±1.9	±1.3	±3.1	±0.0	±14	±23	
	4 mg/kg	N	8	8	8	8	8	8	8	8	8	8	8
		Mean	6.1	0.98	19.9	7.7	17.4	5.6	49.4	0.0	89	59	
		S.D.	±0.2	±0.15	±2.1	±0.9	±2.2	±1.6	±3.8	±0.0	±45	±64	
	20 mg/kg	N	8	8	8	8	8	8	8	8	8	8	8
		Mean	6.2	0.92	18.6	8.1	18.9	7.2	47.2	0.0	93	38	
		S.D.	±0.4	±0.25	±2.3	±1.4	±4.1	±1.8	±6.7	±0.0	±25	±15	
	100 mg/kg	N	10	10	10	10	10	10	10	10	10	10	10
		Mean	6.0	0.79**	18.7	9.2*	22.3**	5.8	44.1**	0.0	101	59*	
		S.D.	±0.4	±0.06	±1.7	±1.1	±1.9	±0.9	±2.1	±0.0	±34	±24	
Female	Control	N	10	10	10	10	10	10	10	10	10	10	
		Mean	6.7	1.49	14.4	5.5	14.6	5.9	59.7	0.1	113	45	
		S.D.	±0.3	±0.16	±1.3	±0.9	±1.5	±0.9	±2.6	±0.1	±69	±56	
	4 mg/kg	N	10	10	10	10	10	10	10	10	10	10	10
		Mean	6.8	1.43	14.4	6.2	14.9	6.0	58.6	0.0	104	32	
		S.D.	±0.2	±0.20	±1.2	±2.3	±2.0	±1.3	±3.6	±0.0	±44	±21	
	20 mg/kg	N	9	9	9	9	9	9	9	9	9	9	9
		Mean	6.9	1.42	14.9	5.6	15.1	5.9	58.5	0.1	86	32	
		S.D.	±0.4	±0.18	±1.5	±0.7	±1.7	±1.0	±3.3	±0.1	±29	±17	
	100 mg/kg	N	10	10	10	10	10	10	10	10	10	10	10
		Mean	7.1*	1.21**	16.5*	5.8	17.3**	5.8	54.5**	0.0	82	31	
		S.D.	±0.4	±0.14	±2.3	±0.6	±2.1	±0.9	±2.9	±0.0	±14	±11	

*: P<0.05, **: P<0.01 (significantly different from control).

One male in the 4 mg/kg group died and one male in the 4 mg/kg group was imminently sacrificed when moribund.

Two males and one female in the 20 mg/kg group died.

Table 9 - continued
Biochemical findings
Male, Female, 52w

Sex	Group and dose		ALP (IU/L)	T.Cholesterol (mg/dL)	Triglycerides (mg/dL)	Phospholipids (mg/dL)	Glucose (mg/dL)	BUN (mg/dL)	Creatinine (mg/dL)	IP (mg/dL)	Ca (mg/dL)	Na (mEq/L)	
Male	Control	N	10	10	10	10	10	10	10	10	10	10	
		Mean	171	85	97	148	121	9.9	0.4	4.9	10.3	146.0	
		S.D.	±51	±11	±37	±21	±17	±1.5	±0.1	±0.4	±0.3	±0.7	
	4 mg/kg	N	8	8	8	8	8	8	8	8	8	8	8
		Mean	150	94	95	157	123	8.7	0.4	4.7	10.0	146.1	
		S.D.	±47	±25	±50	±32	±13	±1.1	±0.1	±0.4	±0.3	±0.8	
	20 mg/kg	N	8	8	8	8	8	8	8	8	8	8	8
		Mean	142	89	93	153	116	9.3	0.4	4.9	10.1	145.6	
		S.D.	±57	±21	±66	±45	±20	±1.7	±0.1	±0.4	±0.3	±0.9	
	100 mg/kg	N	10	10	10	10	10	10	10	10	10	10	10
		Mean	172	76	65	125	128	9.7	0.4	5.1	10.2	146.0	
		S.D.	±63	±15	±30	±20	±19	±1.6	±0.1	±0.9	±0.4	±0.8	
Female	Control	N	10	10	10	10	10	10	10	10	10	10	
		Mean	62	99	52	190	114	13.0	0.5	4.8	10.5	145.4	
		S.D.	±24	±16	±30	±30	±11	±2.1	±0.0	±0.9	±0.4	±0.6	
	4 mg/kg	N	10	10	10	10	10	10	10	10	10	10	10
		Mean	117	106	56	197	105	14.0	0.5	4.7	11.1	145.9	
		S.D.	±235	±11	±26	±20	±15	±4.0	±0.1	±0.5	±1.1	±2.9	
	20 mg/kg	N	9	9	9	9	9	9	9	9	9	9	9
		Mean	60	112	73	212	114	12.8	0.5	4.6	10.9	144.9	
		S.D.	±23	±20	±33	±34	±13	±1.3	±0.1	±0.5	±0.2	±0.6	
	100 mg/kg	N	10	10	10	10	10	10	10	10	10	10	10
		Mean	59	131**	90	228*	116	11.4	0.4	4.8	10.9	144.1**	
		S.D.	±27	±21	±80	±33	±10	±2.8	±0.1	±0.4	±0.3	±0.6	

*: P<0.05, **: P<0.01 (significantly different from control).

One male in the 4 mg/kg group died and one male in the 4 mg/kg group was imminently sacrificed when moribund.

Two males and one female in the 20 mg/kg group died.

Table 9 - continued
Biochemical findings
Male, Female, 52w

Sex	Group and dose		K	Cl
			(mEq/L)	(mEq/L)
Male	Control	N	10	10
		Mean	4.37	105.2
		S.D.	±0.17	±1.6
	4 mg/kg	N	8	8
		Mean	4.30	105.4
		S.D.	±0.16	±1.4
	20 mg/kg	N	8	8
		Mean	4.42	105.7
		S.D.	±0.37	±1.4
	100 mg/kg	N	10	10
		Mean	4.19	104.8
		S.D.	±0.18	±1.4
Female	Control	N	10	10
		Mean	3.91	105.4
		S.D.	±0.29	±2.8
	4 mg/kg	N	10	10
		Mean	3.96	106.5
		S.D.	±0.30	±3.3
	20 mg/kg	N	9	9
		Mean	3.98	105.0
		S.D.	±0.18	±1.4
	100 mg/kg	N	10	10
		Mean	3.99	104.5
		S.D.	±0.27	±1.9

Not significantly different from control.

One male in the 4 mg/kg group died and one male in the 4 mg/kg group was imminently sacrificed when moribund.

Two males and one female in the 20 mg/kg group died.

Table 10 Necropsy findings
Male, Female, 13w

Organs and findings	Sex	Male				Female			
	Group and dose	Control	4 mg/kg	20 mg/kg	100 mg/kg	Control	4 mg/kg	20 mg/kg	100 mg/kg
	Number of animals	10	9	10	9	10	10	10	10
Genital system									
Testis						NA	NA	NA	NA
Enlargement		0	1	0	0	NA	NA	NA	NA
Epididymis									
Nodule, light yellow		0	0	1	0				
Endocrine system									
Pituitary									
Enlargement		0	0	0	0	0	0	0	1
Special sense organs									
Eye									
Dyscoria		1	0	0	0	0	0	0	0

Not significantly different from control.

NA: not applicable.

No appreciable changes in all other organs and tissues.

One male in the 4 mg/kg group was imminently sacrificed when moribund and one male in the 100 mg/kg group died.

Table 11 Necropsy findings
Male, Female, 52w

Organs and findings	Sex	Male				Female			
	Group and dose	Control	4 mg/kg	20 mg/kg	100 mg/kg	Control	4 mg/kg	20 mg/kg	100 mg/kg
	Number of animals	10	8	8	10	10	10	9	10
Digestive system									
Liver									
Macule, dark red		0	0	0	1	1	1	0	0
Enlargement		0	0	1	4	0	0	0	3
Mass, light gray		1	0	0	0	0	0	0	0
Pancreas									
Mass, light gray		0	1	0	0	0	0	0	0
Hematopoietic system									
Thymus									
Small		10	8	8	10	10	10	9	10
Spleen									
Enlargement		0	0	0	1	0	1	0	0
Cyst		0	0	0	0	1	0	0	0
Urinary system									
Kidney									
Rough, surface		0	1	0	0	0	0	0	0
Dilatation, pelvic cavity		0	1	0	0	1	0	0	0
Genital system									
Testis									
Softening		1	0	1	0	NA	NA	NA	NA
Small		0	0	1	0				
Uterus									
Enlargement		NA	NA	NA	NA				
Cyst, endometrium						0	1	0	0
Mammary gland									
Retention, milk		0	1	0	0	7	1*	3	3
Endocrine system									
Pituitary									
Spot, dark red		0	0	1	1	1	1	1	0
Mass, dark red		0	0	1	1	0	1	0	0
Thyroid									
Nodule, light gray		0	0	1	0	0	0	0	0

*: P<0.05 (significantly different from control).

NA: not applicable.

No appreciable changes in all other organs and tissues.

One male in the 4 mg/kg group died and one male in the 4 mg/kg group was imminently sacrificed when moribund.

Two males and one female in the 20 mg/kg group died.

Table 11 - continued

Necropsy findings
Male, Female, 52w

Organs and findings	Sex	Male				Female			
	Group and dose	Control	4 mg/kg	20 mg/kg	100 mg/kg	Control	4 mg/kg	20 mg/kg	100 mg/kg
	Number of animals	10	8	8	10	10	10	9	10
Endocrine system									
Adrenal									
Spot, brown		0	0	0	0	4	1	2	1
Enlargement		0	0	0	0	0	1	0	0
Nodule, brown		0	0	0	0	2	0	0	0
Integumentary system									
Integument									
Mass, subcutis, light gray		1	0	0	0	2	2	2	3
Others									
Extremity									
Swelling, hindlimb		0	0	1	0	0	0	0	1
Corn, hindlimb		4	2	2	4	2	2	1	2

Not significantly different from control.

No appreciable changes in all other organs and tissues.

One male in the 4 mg/kg group died and one male in the 4 mg/kg group was imminently sacrificed when moribund.

Two males and one female in the 20 mg/kg group died.

Table 12 Absolute and relative organ weights
Male, Female, 13w

Sex	Group and dose		Final body weight	Brain		Pituitary		Thyroids		Heart	
			(g)	(g)	(g/100 gB.W.)	(mg)	(mg/100 gB.W.)	(mg)	(mg/100 gB.W.)	(g)	(g/100 gB.W.)
Male	Control	N	10	10	10	10	10	10	10	10	10
		Mean	609.0	2.37	0.39	15.5	2.6	24.6	4.0	1.69	0.28
		S.D.	±49.7	±0.08	±0.03	±2.1	±0.3	±3.4	±0.6	±0.17	±0.02
	4 mg/kg	N	9	9	9	9	9	9	9	9	9
		Mean	582.8	2.36	0.41	14.8	2.6	20.9	3.6	1.66	0.29
		S.D.	±62.6	±0.07	±0.05	±1.8	±0.3	±5.1	±0.9	±0.17	±0.02
	20 mg/kg	N	10	10	10	10	10	10	10	10	10
		Mean	593.6	2.29	0.39	14.9	2.5	26.3	4.4	1.61	0.27
		S.D.	±42.4	±0.12	±0.03	±2.7	±0.3	±4.3	±0.7	±0.15	±0.02
	100 mg/kg	N	9	9	9	9	9	9	9	9	9
		Mean	628.5	2.33	0.37	15.0	2.4	26.4	4.2	1.67	0.27
		S.D.	±37.9	±0.07	±0.02	±2.0	±0.3	±4.5	±0.8	±0.16	±0.02
Female	Control	N	10	10	10	10	10	10	10	10	10
		Mean	316.4	2.05	0.65	17.3	5.6	16.9	5.4	1.02	0.32
		S.D.	±25.1	±0.06	±0.05	±2.7	±1.2	±2.2	±0.8	±0.10	±0.03
	4 mg/kg	N	10	10	10	10	10	10	10	10	10
		Mean	306.7	2.06	0.67	18.2	5.9	16.0	5.3	0.96	0.31
		S.D.	±17.6	±0.07	±0.03	±3.1	±0.9	±2.8	±1.0	±0.07	±0.02
	20 mg/kg	N	10	10	10	10	10	10	10	10	10
		Mean	314.6	2.10	0.67	19.2	6.1	16.8	5.4	1.03	0.33
		S.D.	±34.6	±0.07	±0.06	±3.0	±0.8	±2.5	±0.6	±0.11	±0.02
	100 mg/kg	N	10	10	10	10	10	10	10	10	10
		Mean	307.4	2.13	0.70	19.3	6.3	19.7	6.4*	0.99	0.33
		S.D.	±29.4	±0.06	±0.07	±4.0	±1.0	±3.9	±1.1	±0.09	±0.03

*: P<0.05 (significantly different from control).

One male in the 4 mg/kg group was imminently sacrificed when moribund and one male in the 100 mg/kg group died.

Table 12 - continued Absolute and relative organ weights
Male, Female, 13w

Sex	Group and dose		Lungs		Thymus		Liver		Spleen	
			(g)	(g/100 gB.W.)	(g)	(g/100 gB.W.)	(g)	(g/100 gB.W.)	(g)	(g/100 gB.W.)
Male	Control	N	10	10	10	10	10	10	10	10
		Mean	1.71	0.28	0.31	0.05	16.72	2.75	0.91	0.15
		S.D.	±0.10	±0.02	±0.10	±0.02	±1.53	±0.13	±0.13	±0.01
	4 mg/kg	N	9	9	9	9	9	9	9	9
		Mean	1.64	0.28	0.33	0.06	16.25	2.79	0.87	0.15
		S.D.	±0.15	±0.02	±0.09	±0.02	±2.42	±0.23	±0.10	±0.02
	20 mg/kg	N	10	10	10	10	10	10	10	10
		Mean	1.63	0.28	0.30	0.05	16.94	2.85	0.81	0.14
		S.D.	±0.14	±0.03	±0.10	±0.02	±1.83	±0.21	±0.12	±0.01
	100 mg/kg	N	9	9	9	9	9	9	9	9
		Mean	1.72	0.28	0.28	0.05	20.20**	3.21**	0.88	0.14
		S.D.	±0.10	±0.02	±0.07	±0.01	±1.76	±0.20	±0.14	±0.02
Female	Control	N	10	10	10	10	10	10	10	10
		Mean	1.14	0.36	0.27	0.09	8.24	2.61	0.51	0.16
		S.D.	±0.07	±0.02	±0.06	±0.02	±0.81	±0.20	±0.05	±0.02
	4 mg/kg	N	10	10	10	10	10	10	10	10
		Mean	1.19	0.39	0.27	0.09	8.42	2.75	0.50	0.16
		S.D.	±0.07	±0.03	±0.06	±0.02	±0.65	±0.19	±0.07	±0.02
	20 mg/kg	N	10	10	10	10	10	10	10	10
		Mean	1.22	0.39	0.27	0.08	9.24	2.93**	0.55	0.18
		S.D.	±0.10	±0.04	±0.06	±0.02	±1.40	±0.21	±0.07	±0.02
	100 mg/kg	N	10	10	10	10	10	10	10	10
		Mean	1.22	0.40*	0.28	0.09	10.51**	3.42**	0.51	0.17
		S.D.	±0.09	±0.03	±0.08	±0.03	±1.18	±0.23	±0.06	±0.02

*: P<0.05, **: P<0.01 (significantly different from control).

One male in the 4 mg/kg group was imminently sacrificed when moribund and one male in the 100 mg/kg group died.